

## Complete Summary

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### GUIDELINE TITLE

Unstable angina pectoris.

### BIBLIOGRAPHIC SOURCE(S)

Finnish Medical Society Duodecim. Unstable angina pectoris. In: EBM Guidelines. Evidence-Based Medicine [CD-ROM]. Helsinki, Finland: Duodecim Medical Publications Ltd.; 2004 Sep 14 [Various].

### GUIDELINE STATUS

This is the current release of the guideline.

This guideline updates a previous version: Finnish Medical Society Duodecim. Unstable angina pectoris. Helsinki, Finland: Duodecim Medical Publications Ltd.; 2003 Jul 11. Various p.

## COMPLETE SUMMARY CONTENT

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## SCOPE

### DISEASE/CONDITION(S)

Unstable angina pectoris

### GUIDELINE CATEGORY

Diagnosis  
 Evaluation  
 Risk Assessment  
 Treatment

### CLINICAL SPECIALTY

Cardiology  
Family Practice  
Internal Medicine

## INTENDED USERS

Health Care Providers  
Physicians

## GUIDELINE OBJECTIVE(S)

Evidence-Based Medicine Guidelines collects, summarizes, and updates the core clinical knowledge essential in general practice. The guidelines also describe the scientific evidence underlying the given recommendations.

## TARGET POPULATION

Patients with suspected or confirmed unstable angina

## INTERVENTIONS AND PRACTICES CONSIDERED

### Diagnosis

1. Electrocardiogram
2. Assessment of signs and symptoms
3. Measurement of myocardial markers, such as troponin T or troponin I and creatine kinase isoenzyme MB mass (CK-MBm)

### Treatment

1. Aspirin
2. Oxygen
3. Nitrate infusion
4. Beta-blockers, such as metoprolol or atenolol
5. Low-molecular-weight (LMW) heparins (such as dalteparin, nadroparin, enoxaparin) simultaneously with aspirin
6. Cardiac monitoring
7. Angiography
8. Revascularization
9. Clopidogrel and intravenous glycoprotein IIb/IIIa (GP IIb/IIIa) inhibitor in addition to aspirin and low-molecular-weight heparin
10. Thrombolytic therapy
11. Percutaneous transluminal angioplasty (PTA) with insertion of a stent followed by clopidogrel in combination with aspirin
12. Elimination of risk factors

### Evaluation

1. Electrocardiogram
2. Measurement of myocardial markers, such as troponin T or troponin I and creatine kinase isoenzyme MB mass

### 3. Exercise testing

#### MAJOR OUTCOMES CONSIDERED

- Mortality
- Incidence of myocardial infarction
- Efficacy of treatment for angina in reducing mortality and incidence of myocardial infarction
- Episodes of chest pain and severity of chest pain (i.e., need for sublingual nitroglycerin)
- Incidence of ischaemic events/recurrent angina
- Bleeding complications
- Need for urgent revascularization
- Sensitivity and specificity of troponin I and T for predicting adverse cardiac events in unstable angina pectoris

## METHODOLOGY

#### METHODS USED TO COLLECT/SELECT EVIDENCE

Hand-searches of Published Literature (Primary Sources)  
Hand-searches of Published Literature (Secondary Sources)  
Searches of Electronic Databases

#### DESCRIPTION OF METHODS USED TO COLLECT/SELECT THE EVIDENCE

The evidence reviewed was collected from the Cochrane database of systematic reviews and the database of abstracts of reviews of effectiveness (DARE). In addition, the Cochrane Library and medical journals were searched specifically for original publications.

#### NUMBER OF SOURCE DOCUMENTS

Not stated

#### METHODS USED TO ASSESS THE QUALITY AND STRENGTH OF THE EVIDENCE

Weighting According to a Rating Scheme (Scheme Given)

#### RATING SCHEME FOR THE STRENGTH OF THE EVIDENCE

##### Levels of Evidence

- A. Strong research-based evidence. Multiple relevant, high-quality scientific studies with homogenic results.
- B. Moderate research-based evidence. At least one relevant, high-quality study or multiple adequate studies.
- C. Limited research-based evidence. At least one adequate scientific study.
- D. No research-based evidence. Expert panel evaluation of other information.

## METHODS USED TO ANALYZE THE EVIDENCE

Review of Published Meta-Analyses  
Systematic Review

## DESCRIPTION OF THE METHODS USED TO ANALYZE THE EVIDENCE

Not stated

## METHODS USED TO FORMULATE THE RECOMMENDATIONS

Not stated

## RATING SCHEME FOR THE STRENGTH OF THE RECOMMENDATIONS

Not applicable

## COST ANALYSIS

A formal cost analysis was not performed and published cost analyses were not reviewed.

## METHOD OF GUIDELINE VALIDATION

Peer Review

## DESCRIPTION OF METHOD OF GUIDELINE VALIDATION

Not stated

# RECOMMENDATIONS

## MAJOR RECOMMENDATIONS

The levels of evidence [A-D] supporting the recommendations are defined at the end of the "Major Recommendations" field.

### Objective

- To recognize angina pectoris that may be prodromal to acute infarction (acute coronary syndrome [ACS]) and to accompany the patient to a cardiac monitoring unit for active drug treatment or rapid revascularization (Wallentin et al., 2000) [B].

### Definition

- Recent (less than 1 to 2 months) angina pectoris
- Accelerated angina pectoris
- Angina pectoris at rest

## Risk Groups and Clinical Signs

- The presence of a marker (cardiac troponin T and I, creatine kinase isoenzyme MB mass [CK-MBm]) is the single most important predictor for future coronary events.
  - Marker-positive patients are referred to angiography and revascularization.
  - Marker-negatives are referred to exercise tolerance test.
- Unstable angina pectoris (UAP) is a heterogeneous group of diseases covering the range between stable angina pectoris (AP) and acute myocardial infarction (AMI).
- New (sudden) AP in a high-risk patient is always a serious condition.
- An aggravation in stable AP to unstable AP always necessitates a reassessment of risk and often a change in the line of treatment.
- There may not always be pain; rather, the main symptom is a decrease in exercise tolerance (sudden decrease in physical fitness) or acute left ventricle failure.
- In the electrocardiogram (ECG) an ST segment depression precedes the pain. Symptomless (silent) ischaemia in a patient at risk is a significant finding. Ischaemia may not always be visible in ECG. An ECG registered while the patient has pain is invariably valuable.
- The border between unstable AP and T-wave infarction (non-Q infarction) is shifting. For example, very proximal occlusion in the left anterior descending artery (LAD) causes a symmetric T inversion in chest leads. Elevation of myocardial markers indicates that the patient has an infarction.

## Treatment

- Treatment is normally carried out in a cardiac monitoring unit.
- Pharmacological treatment should be started in the first point of care.
- The mildest form (recent angina) can be treated in a health care centre ward under careful monitoring. Remember the risk of myocardial infarction (MI). The risk diminishes with time as the angina stabilizes.

## Anti-ischaemic and Antithrombotic Treatment

- All patients with suspected unstable angina (no changes in ECG or myocardial markers)
  - Aspirin 250 mg (chewable) first. Thereafter 100 mg/day, unless there are contraindications (Natarajan, 2002; "Collaborative overview," 1994) [A].
  - Oxygen
  - Nitrate infusion (Natarajan, 2000) [D] for 24 to 36 hours (see the related Evidence-Based Medicine [EBM] guideline "Nitrate Infusion in Angina Pectoris and Myocardial Infarction"). Systolic blood pressure should be lowered by 10 to 15 mmHg and always to a level below 150 mmHg.
  - Beta-blocker (Natarajan, 2002) [C] (metoprolol or atenolol). Heart rate should be 50 to 70 beats per minute and systolic pressure below 150 mmHg.
  - Low-molecular-weight (LMW) heparin (Zed, Tisdale, & Borzak, 1999; DARE-999714, 2001; Nicholson, Milne, & Stein, 2000; The Health

Technology Assessment Database, HTA-20000891, 2001) [A] (e.g., dalteparin 100-120 IU x 2 daily for one week) is given simultaneously with aspirin. The treatment can be continued with half the dose for about 1 month. Unstable AP patients with an elevated troponin T concentration derive the greatest benefit from the treatment (low-molecular-weight heparin + aspirin). Pharmacotherapy and invasive treatment do not exclude one another.

- High-risk patients: Unstable angina and ischaemia on ECG or elevated myocardial markers (Olatidoye et al., 1998; DARE-981100, 2000) [A], acute left ventricle failure (lung oedema, mitral regurgitation, hypotension)
  - Immediate angiography and revascularization. While waiting for the procedure, the thrombosis should be stabilized with clopidogrel (an initial dose of 300 mg before transportation, thereafter 75 mg daily) and an intravenous (i.v.) glycoprotein IIb/IIIa (GP IIb/IIIa) inhibitor (Natarajan, 2002) [B] in addition to aspirin and low-molecular-weight heparin. (Fibrinolytic treatment has no effect on a vessel obstruction caused by aggregated platelets.)
- Thrombolytic therapy or immediate percutaneous transluminal angioplasty (PTA) (during which a stent can be inserted) is indicated if ECG reveals a transmural injury. (See article on revascularization: Evidence-Based Medicine guideline, "Coronary heart disease symptoms, diagnosis and treatment.") After the insertion of a stent, clopidogrel is used in combination with aspirin for one month.
- Further treatment of patients with symptoms or signs of ischaemia on ECG and normal myocardial markers
  - Symptom-limited exercise test performed within 2 to 4 days.
  - If the patient has symptoms or signs of ischaemia during the exercise test or signs in ECG at a low pulse-pressure product, refer immediately to angiography.
  - In case of no symptoms or signs of ischaemia during light exercise or no signs in ECG, or if they occur only with a high pulse-pressure product, begin conservative treatment and elimination of risk factors. Prophylaxis can be intensified by adding clopidogrel to aspirin.

## Organizing Treatment

- Unstable AP is a serious but often curable syndrome. A well-organized care pathway ensures that the appropriate treatment can be given rapidly.

## Related Evidence

- Intravenous heparin combined with aspirin is probably effective compared to aspirin alone in reducing myocardial infarction or death, but the number of patients studied is too small for statistical significance (Oler et al., 1996; DARE-968398, 1999) [C].

## Definitions:

## Levels of Evidence

- A. Strong research-based evidence. Multiple relevant, high-quality scientific studies with homogenic results.

- B. Moderate research-based evidence. At least one relevant, high-quality study or multiple adequate studies.
- C. Limited research-based evidence. At least one adequate scientific study.
- D. No research-based evidence. Expert panel evaluation of other information.

#### CLINICAL ALGORITHM(S)

None provided

### EVIDENCE SUPPORTING THE RECOMMENDATIONS

#### REFERENCES SUPPORTING THE RECOMMENDATIONS

[References open in a new window](#)

#### TYPE OF EVIDENCE SUPPORTING THE RECOMMENDATIONS

Concise summaries of scientific evidence attached to the individual guidelines are the unique feature of the Evidence-Based Medicine Guidelines. The evidence summaries allow the clinician to judge how well-founded the treatment recommendations are. The type of supporting evidence is identified and graded for select recommendations (see the "Major Recommendations" field).

### BENEFITS/HARMS OF IMPLEMENTING THE GUIDELINE RECOMMENDATIONS

#### POTENTIAL BENEFITS

##### Overall Benefits

- Appropriate diagnosis and treatment of unstable angina pectoris
- Reduced rate of myocardial infarction and death

##### Specific Benefits

- An invasive revascularization strategy improves survival in patients with unstable coronary artery disease.
- There is no good evidence on the effect of nitrates on prevention of death or myocardial infarction in patients with unstable angina. Still, nitrates are effective in pain relief and remain first-line treatment together with heparin and aspirin in unstable angina.
- Low-molecular-weight heparins are superior to placebo and unfractionated heparin in reducing ischaemic events or death in acute unstable angina or non-Q-wave myocardial infarction.
- Troponin T and troponin I predict adverse cardiac events in patients with unstable angina pectoris.
- Intravenous heparin combined with aspirin is probably effective compared to aspirin alone in reducing myocardial infarction or death, but the number of patients studied is too small for statistical significance.

#### POTENTIAL HARMS

Low-molecular-weight heparin can cause bleeding complications.

## IMPLEMENTATION OF THE GUIDELINE

### DESCRIPTION OF IMPLEMENTATION STRATEGY

An implementation strategy was not provided.

## INSTITUTE OF MEDICINE (IOM) NATIONAL HEALTHCARE QUALITY REPORT CATEGORIES

### IOM CARE NEED

Getting Better  
Staying Healthy

### IOM DOMAIN

Effectiveness  
Timeliness

## IDENTIFYING INFORMATION AND AVAILABILITY

### BIBLIOGRAPHIC SOURCE(S)

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### ADAPTATION

Not applicable: The guideline was not adapted from another source.

### DATE RELEASED

2001 Apr 30 (revised 2004 Sep 14)

### GUIDELINE DEVELOPER(S)

Finnish Medical Society Duodecim - Professional Association

### SOURCE(S) OF FUNDING

Finnish Medical Society Duodecim

### GUIDELINE COMMITTEE

Editorial Team of EBM Guidelines

## COMPOSITION OF GROUP THAT AUTHORED THE GUIDELINE

Primary Authors: Editors

## FINANCIAL DISCLOSURES/CONFLICTS OF INTEREST

Not stated

## GUIDELINE STATUS

This is the current release of the guideline.

This guideline updates a previous version: Finnish Medical Society Duodecim. Unstable angina pectoris. Helsinki, Finland: Duodecim Medical Publications Ltd.; 2003 Jul 11. Various p.

## GUIDELINE AVAILABILITY

This guideline is included in a CD-ROM titled "EBM Guidelines. Evidence-Based Medicine" available from Duodecim Medical Publications, Ltd, PO Box 713, 00101 Helsinki, Finland; e-mail: [info@ebm-guidelines.com](mailto:info@ebm-guidelines.com); Web site: [www.ebm-guidelines.com](http://www.ebm-guidelines.com).

## AVAILABILITY OF COMPANION DOCUMENTS

None available

## PATIENT RESOURCES

None available

## NGC STATUS

This summary was completed by ECRI on August 28, 2001. The information was verified by the guideline developer as of October 26, 2001. This summary was updated by ECRI on December 9, 2002, April 2, 2004, and February 22, 2005.

## COPYRIGHT STATEMENT

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